

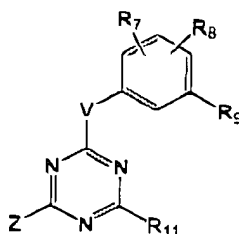
US application Serial No. 09/891/750  
 Attorney Docket No. QA0239A-CIP

AMENDMENTS:

Cancel claims 52-65, 67-69, 83, 84, 86, 87, 89 - 95

Replace claims 66 and 70-95 as follows:

66 (Amended). A compound of Formula (I),



I

or an enantiomer, diastereomer, tautomer, or pharmaceutically-acceptable salt, or solvate thereof, wherein:

V is chosen from  $-\text{CHR}^5-$ ,  $-\text{NR}^5-$ ,  $-\text{O}-$ , and  $-\text{S}-$ ;

Z is chosen from halogen, alkyl, substituted alkyl, aryl, substituted aryl, cycloalkyl, substituted cycloalkyl, heterocyclyl, substituted heterocyclyl,  $-\text{SR}^3$ ,  $-\text{OR}^3$ , and  $-\text{N}(\text{R}^1)(\text{R}^2)$ ;  $-\text{N}(\text{R}^1)(\text{R}^2)$  taken together may form a heterocyclyl or substituted heterocyclyl; or

$\text{R}^1$  is chosen from hydrogen, alkyl and substituted alkyl; and

$\text{R}^2$  is chosen from hydrogen, alkyl, substituted alkyl, alkoxy, aryl, substituted aryl, cycloalkyl, substituted cycloalkyl, heterocyclyl and substituted heterocyclyl;

$\text{R}^3$  is chosen from hydrogen, alkyl, substituted alkyl, aryl, substituted aryl, cycloalkyl, substituted cycloalkyl, heterocyclyl and substituted heterocyclyl;

$\text{R}^5$  is chosen from hydrogen and alkyl, or when attached to a nitrogen atom,  $\text{R}^5$  taken together with  $\text{R}^7$  may form a fused heterocyclyl or substituted heterocyclyl;

$\text{R}^7$  is chosen from hydrogen,  $-\text{N}(\text{R}^{31})(\text{R}^{32})$ , halogen, cyano, alkyl, substituted alkyl, alkoxy, and alkylthio, or when V is  $-\text{NR}^5$ ,  $-\text{R}^5$  and  $\text{R}^7$  taken together may form a fused heterocyclyl or substituted heterocyclyl;

$\text{R}^8$  is chosen from hydrogen and halogen;

$\text{R}^9$  is chosen from  $-\text{CO}_2(\text{alkyl})$ ,  $-\text{C}(\text{O})\text{N}(\text{R}^{31})(\text{R}^{32})$ ,  $-\text{SO}_2\text{N}(\text{R}^{31})(\text{R}^{32})$ ,

US application Serial No. 09/891/750  
 Attorney Docket No. QA0239A-CIP

$-N(R^{33})SO_2R^{34}$ ,  $-C(O)N(R^{33})N(R^{31})(R^{32})$ ,  $-N(R^{33})C(O)R^{34}$ ,  $-CH_2N(R^{33})C(O)R^{34}$ ,  $-N(R^{31})(R^{32})$ ,  $-CH_2OC(O)R^{34}$ ,  $C_{1-6}$ alkyl, substituted alkyl, cycloalkyl, substituted cycloalkyl, aryl, substituted aryl, heterocyclyl, substituted heterocyclyl, and  $-C(O)R^{10}$ ; provided, however, that when  $R^9$  is  $CH_3$  or  $NH_2$ , then neither  $R^2$  nor  $R^{14}$  is *para*-cyano-phenyl;

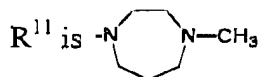
or  $R^8$  and  $R^9$  taken together may form  $-C(O)N(R^{33})CH_2-$  or  $-C(O)N(R^{33})C(O)-$ ;

$R^{10}$  is chosen from heterocyclyl, substituted heterocyclyl, cycloalkyl, substituted cycloalkyl, aryl, substituted aryl, alkyl, and substituted alkyl;

$R^{31}$  and  $R^{33}$  are independently chosen from hydrogen, alkyl, and substituted alkyl;

$R^{32}$  is chosen from hydrogen, alkyl, substituted alkyl, alkoxy, aryl, substituted aryl, cycloalkyl, substituted cycloalkyl, aryloxy, heterocyclyl and substituted heterocyclyl;

$R^{34}$  is chosen from alkyl, substituted alkyl, aryl, substituted aryl, cycloalkyl, substituted cycloalkyl, heterocyclyl and substituted heterocyclyl;



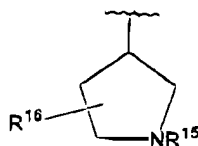
$R^{12}$  is chosen from hydrogen, alkyl, and substituted alkyl;

$R^{13}$  is  $-(CH_2)_mR^{14}$ ;

$-N(R^{12})(R^{13})$  taken together may form a heterocyclyl or substituted heterocyclyl;

$m$  is 0, 1, 2 or 3;

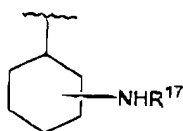
$R^{14}$  is chosen from hydrogen, alkyl, substituted alkyl,  $-C(O)N(R^{31})(R^{32})$ ,  $-N(R^{33})C(O)R^{34}$ , aryl, substituted aryl, cycloalkyl, substituted cycloalkyl, heterocyclyl, substituted heterocyclyl, and



$R^{15}$  is chosen from hydrogen, alkyl, substituted alkyl, alkenyl,  $-C(O)$ -alkyl,  $-C(O)$ -substituted alkyl,  $-C(O)$ -aryl,  $-C(O)$ -substituted aryl,  $-C(O)$ -alkoxy, aryl, substituted aryl, cycloalkyl, substituted cycloalkyl, heterocyclyl and substituted heterocyclyl;

$R^{16}$  is chosen hydrogen, alkyl, substituted alkyl, and

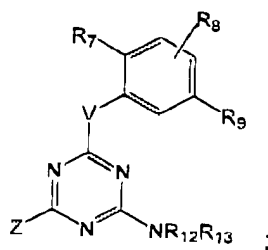
US application Serial No. 09/891/750  
 Attorney Docket No. QA0239A-CIP



or

$R^{17}$  is chosen from hydrogen, alkyl, substituted alkyl,  $-C(O)$ -alkyl,  $-C(O)$ -substituted alkyl,  $-C(O)$ -aryl, and  $-C(O)$ -substituted aryl.

70. (Amended). A compound having the formula,



or a enantiomer, diastereomer, tautomer, or pharmaceutically-acceptable salt, or solvate thereof, wherein:

V is chosen from  $-CHR^5$ -,  $-NR^5$ -,  $-O$ -, and  $-S$ -;

Z is halogen, alkyl,  $-N(R^1)(R^2)$ , or alkyl substituted with one to two of  $-N(R^{31})(R^{32})$ , alkoxy, alkylthio, halogen, cyano, carboxyl, hydroxyl,  $-SO_2$ -alkyl,  $-CO_2$ -alkyl,  $-C(O)$ -alkyl, nitro, cycloalkyl, substituted cycloalkyl,  $-C(O)-N(R^{31})(R^{32})$ , and/or  $-NH-C(O)$ -alkyl;

$R^1$  is hydrogen or methyl;

$R^2$  is alkyl of 1 to 8 carbon atoms;

$R^3$  is chosen from hydrogen, alkyl, substituted alkyl, aryl, substituted aryl, cycloalkyl, substituted cycloalkyl, heterocyclyl and substituted heterocyclyl;

$R^5$  is chosen from hydrogen and alkyl of 1 to 4 carbon atoms;

$R^7$  is chosen from hydrogen, amino, amino $C_{1-4}$ alkyl, halogen, cyano,  $C_{1-4}$ alkyl,  $C_{1-4}$ alkoxy, and alkylthio;

$R^8$  is attached to any available carbon atom of the phenyl ring and is chosen from hydrogen and halogen;

US application Serial No. 09/891/750

Attorney Docket No. QA0239A-CIP

$R^9$  is chosen from  $-C(O)N(R^{31})(R^{32})$ ,  $-SO_2N(R^{31})(R^{32})$ ,  $-N(R^{33})SO_2R^{34}$ ,  $-C(O)N(R^{33})N(R^{31})(R^{32})$ ,  $-N(R^{33})C(O)R^{34}$ ,  $-CH_2N(R^{33})C(O)R^{34}$ ,  $-N(R^{31})(R^{32})$ ,  $-CH_2OC(O)R^{34}$ , heterocyclyl, and substituted heterocyclyl; or

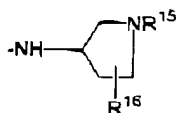
$R^8$  and  $R^9$  taken together may form  $-C(O)N(R^{33})CH_2-$  or  $-C(O)N(R^{33})C(O)-$ ;

$R^{31}$  and  $R^{33}$  are independently chosen from hydrogen, alkyl, and substituted alkyl;

$R^{32}$  is chosen from hydrogen, alkyl, substituted alkyl, alkoxy, aryl, substituted aryl, cycloalkyl, substituted cycloalkyl, aryloxy, heterocyclyl and substituted heterocyclyl;

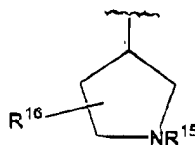
$R^{34}$  is chosen from alkyl, substituted alkyl, aryl, substituted aryl, cycloalkyl, substituted cycloalkyl, heterocyclyl and substituted heterocyclyl;

$N(R^{12})(R^{13})$  taken together form a monocyclic heterocyclyl or substituted heterocyclyl of 5 to 7 atoms having 1, 2 or 3 additional nitrogen atoms,  $-NH$ -alkyl wherein alkyl is of 1 to 4 carbon atoms, or



$m$  is 0, 1, 2 or 3;

$R^{14}$  is chosen from hydrogen, alkyl, substituted alkyl,  $-C(O)N(R^{31})(R^{32})$ ,  $-N(R^{33})C(O)R^{34}$ , aryl, substituted aryl, cycloalkyl, substituted cycloalkyl, heterocyclyl, substituted heterocyclyl and

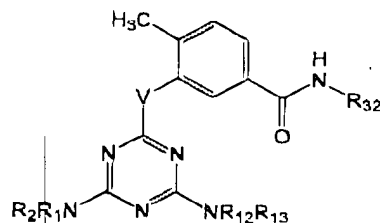


$R^{15}$  and  $R^{16}$  are independently hydrogen or methyl; and

$R^{17}$  is chosen from hydrogen, alkyl, substituted alkyl,  $-C(O)$ -alkyl,  $-C(O)$ -substituted alkyl,  $-C(O)$ -aryl, and  $-C(O)$ -substituted aryl.

71 (Amended). A compound of Claim 70 or a enantiomer, diastereomer, tautomer, or pharmaceutically-acceptable salt, or solvate thereof, having the formula:

US application Serial No. 09/891/750  
 Attorney Docket No. QA0239A-CIP



72 (Amended). The compound of claim 70 or a enantiomer, diastereomer, tautomer, or pharmaceutically-acceptable salt, or solvate thereof, wherein:

$R^7$  is halogen, methyl, methoxy, halogen, or cyano.

73 (Amended). The compound of claim 70 or a stereoisomer, enantiomer, diastereomer, tautomer, or pharmaceutically-acceptable salt, prodrug, or solvate thereof, wherein:

$R^9$  is  $C(=O)NH_2$ ,  $C(=O)NH(CH_3)$ , or  $C(=O)NHO(CH_3)$ .

74 (Amended). The compound of claim 70 or a enantiomer, diastereomer, tautomer, or pharmaceutically-acceptable salt, or solvate thereof,

wherein  $R^7$  is methyl and  $R^9$  is  $C(=O)NH(CH_3)$  or  $C(=O)NHO(CH_3)$ .

75 (Amended). A compound of Claim 70 or a enantiomer, diastereomer, tautomer, or pharmaceutically-acceptable salt, or solvate thereof wherein:

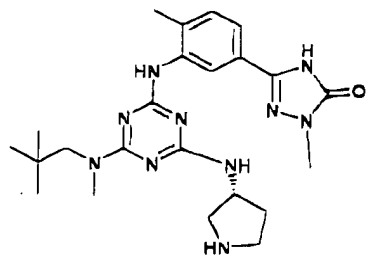
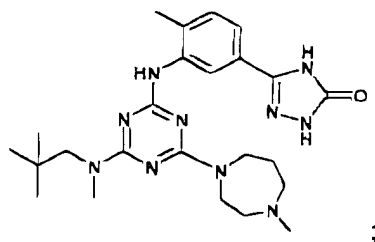
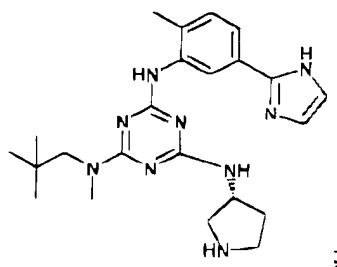
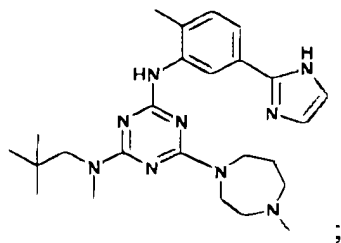
$R^9$  is chosen from unsubstituted or substituted triazolyl, oxadiazolyl, imidazolyl, thiazolyl and benzimidazolyl.

76 (Amended). A compound of Claim 70 or a enantiomer, diastereomer, tautomer, or pharmaceutically-acceptable salt, or solvate thereof  
 wherein:

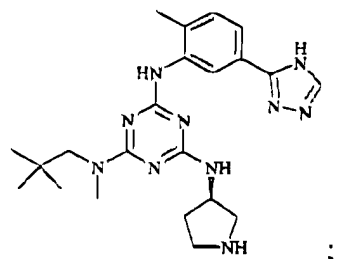
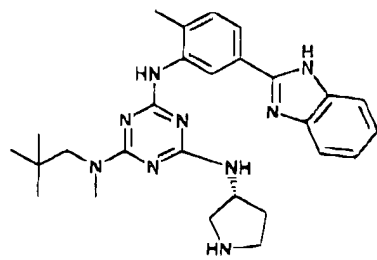
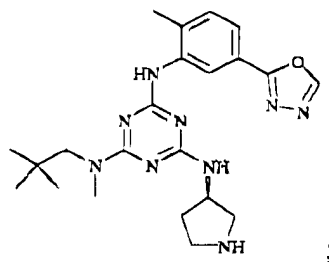
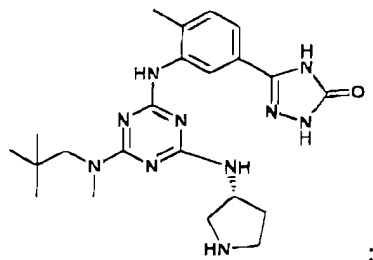
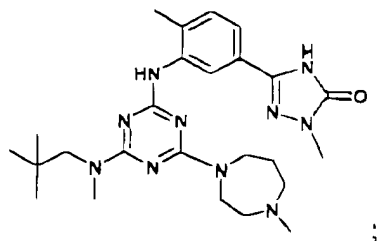
$R^9$  is chosen from substituted or unsubstituted 1,2,4-triazole; substituted or unsubstituted thiazole connected via a C2, C4, or C5 position; substituted or unsubstituted 1,3,4-oxdiazole connected via a 2 or 5 position; and substituted or unsubstituted imidazole connected via a C2, C4, C5, N1 or N3 position.

US application Serial No. 09/891/750  
Attorney Docket No. QA0239A-CIP

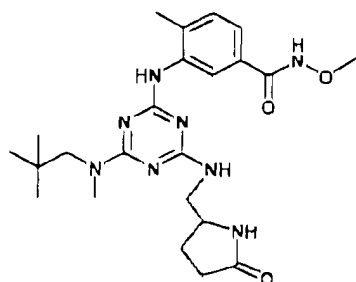
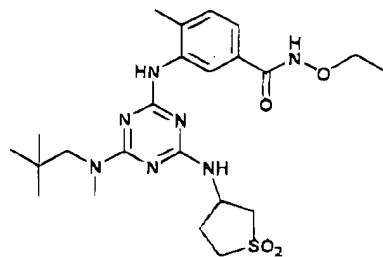
77 (Amended). A compound which is selected from (i):



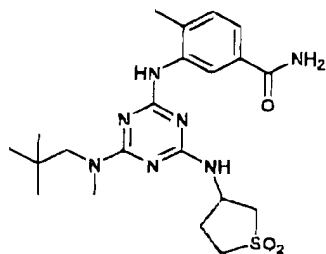
US application Serial No. 09/891/750  
Attorney Docket No. QA0239A-CIP



US application Serial No. 09/891/750  
Attorney Docket No. QA0239A-CIP



; and



; or (ii) a enantiomer, diastereomer, tautomer, or pharmaceutically-acceptable salt, or solvate of the compound selected from paragraph (i).

78 (Amended). A pharmaceutical composition comprising as an active ingredient, a compound, or a prodrug or salt thereof, according to claim 70, and a pharmaceutically acceptable carrier.

82 (New). A method of treating rheumatoid arthritis, the method comprising administering to a mammal in need of such treatment, an effective amount of a composition according to claim 78.